DO 3/2/05 MANY

Attorney Docket No.: 23546-08263

Client Ref.: ISIS-2960

Application No.: 09/067,638

## AMENDMENTS TO THE CLAIMS

1-82. (Canceled)

83. (Currently amended) A system of associated components for preparing a set of oligonucleotides that modulates expression of a selected nucleic acid comprising:

a computer system that prepares a virtual library of oligonucleotide[s] sequences targeted to said selected nucleic acid and generates synthesis instructions in computer manipulable form for [each of ]said oligonucleotide[s] sequences in said virtual library, wherein said computer system first prepares said virtual library of oligonucleotide sequences and then reduces the number of oligonucleotide sequences in [members of ]said virtual library of oligonucleotide[s] sequences by a process of selection based on targeting a functional region of said selected nucleic acid;

an automated synthesizer that receives said synthesis instructions from said computer system and synthesizes only that [a ]set of real oligonucleotides that corresponds [corresponding] to said virtual set of oligonucleotide[s] sequences [having] consisting of said reduced number of oligonucleotide sequences [members]; and

an apparatus that accepts said set of real oligonucleotides and performs at least one procedure for each of said real oligonucleotides wherein said procedure identifies particular members of said set that modulate expression of said selected nucleic acid, wherein said procedure is computer-controlled polymerase chain reaction or computer-controlled enzymelinked immunosorbent assay.

84. (Previously presented) The system of claim 83 wherein said functional region is the transcription start site, 5' cap, start codon, 5' untranslated region, 3' untranslated region, stop codon, 5' splice site or polyadenylation site.